



RECEIVED

APR 23 2001

PATENT

TECH CENTER 1600/2900

16  
S. H. 16  
#14/C

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants: Gruber, et al.

Examiner: Bui, P.

Serial No.: 09/284,697

Group Art Unit: 1638

Filed: July 6, 1999

Docket: 1149-2

For: PANCREATIC LIPASES  
AND/OR RECOMBINANT  
COLIPASES AND DERIVED  
POLYPEPTIDES PRODUCED  
BY PLANTS, METHODS FOR  
OBTAINING THEM AND USE  
THEREOF

Dated: April 12, 2001

I hereby certify this correspondence is being deposited with the United States Postal Service as first class mail, postpaid in an envelope, addressed to:  
Assistant Commissioner for Patents, Washington, D.C.

20231 on April 12, 2001  
Dated: 4/12/01 J. C. L.

Assistant Commissioner for Patents  
Washington, DC 20231

REQUEST FOR ENTRY OF THE REVISED SEQUENCE LISTING

Sir:

This Request is **not** a response to the Office Action that was issued along with the Raw Sequence Listing Error Report. A response to the Office Action will follow.

Along with this Request, Applicants submit the following:

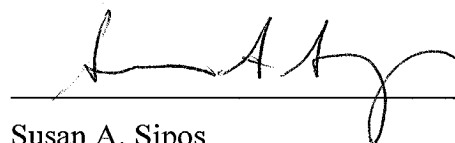
1. An substitute Diskette containing the revised Sequence Listing in computer-readable form;
2. An substitute paper copy of the revised Sequence Listing;
3. A copy of the Raw Sequence Listing Error Report; and
4. Response to Raw Sequence Listing Error Report.

Entry of the revised Sequence Listing into the specification of the application is respectfully requested.

The revised Sequence Listing contains no new matter. Only the formatting changes specified in the Error Report were made to the original Sequence Listing. The contents of the substitute computer readable form and the substitute paper copy are the same.

If the Examiner has any questions relating to this application, it is respectfully requested that the Examiner contact the Applicants' undersigned attorney at the telephone number provided below.

Respectfully submitted,

A handwritten signature in black ink, appearing to read 'Susan A. Sipos', is written over a horizontal line.

Susan A. Sipos  
Registration No. 43,128  
Attorney for Applicants

HOFMANN & BARON, LLP  
6900 Jericho Turnpike  
Syosset, New York 11791  
(516) 822-3550  
SAS/dlv/jjc

131641\_1

PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants: Gruber, et al.

Examiner: Bui, P.

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For: PANCREATIC LIPASES  
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COLIPASES AND DERIVED  
POLYPEPTIDES PRODUCED  
BY PLANTS, METHODS FOR  
OBTAINING THEM AND USE  
THEREOF

Dated: April 12, 2001

I hereby certify this correspondence is being deposited with the United States Postal Service as first class mail, postpaid in an envelope, addressed to: Assistant Commissioner for Patents, Washington, D.C.

20231 on

Dated:

Assistant Commissioner for Patents  
Washington, DC 20231

RESPONSE TO RAW SEQUENCE LISTING ERROR REPORT

Sir:

This Response is **not** a response to Office Action that was issued along with the Raw Sequence Listing Error Report. A response to the Office Action will follow. The present Response only addresses the Raw Sequence Listing Error Report mailed March 22, 2001.

Applicants enclose herewith the following:

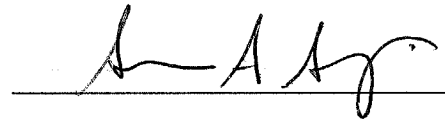
1. An substitute Diskette containing the revised Sequence Listing in computer-readable form;
2. An substitute paper copy of the revised Sequence Listing;
3. A copy of the Raw Sequence Listing Error Report; and
4. Request for Entry of the Revised Sequence Listing.

Applicants submit that the substitute Sequence Listing complies with the requirements set forth in the Raw Sequence Listing Error Report.

Additionally, under 37 C.F.R. 1.821(f), the Applicants' attorney hereby states that the contents of the substitute computer readable form and the substitute paper copy are the same. Under 37 C.F.R. §1.821(g), the Applicants' attorney also states that this Sequence Listing does not include any new matter.

If the Examiner has any questions relating to this application, it is respectfully requested that the Examiner contact the Applicants' undersigned attorney at the telephone number provided below.

Respectfully submitted,

A handwritten signature in black ink, appearing to read 'S. A. Sipos', is written over a horizontal line.

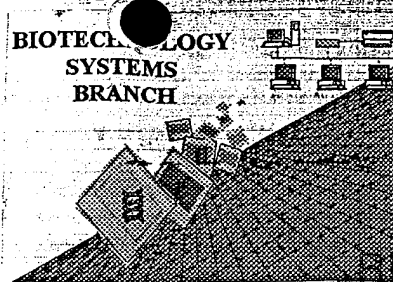
Susan A. Sipos  
Registration No. 43,128  
Attorney for Applicants

HOFMANN & BARON, LLP  
6900 Jericho Turnpike  
Syosset, New York 11791  
(516) 822-3550  
SAS/dlv

2/Bur

## RAW SEQUENCE LISTING ERROR REPORT

BIOTECHNOLOGY  
SYSTEMS  
BRANCH



RECEIVED

The Biotechnology Systems Branch of the Scientific and Technical Information Center (STIC) detected errors when processing the following computer readable form: DEC 14 2000

TECH CENTER 1600/2900

Application Serial Number: 09/284,697

Source: 1638

Date Processed by STIC: 12/1/2000

Lib 6573  
mwl 703-308-4266  
Spencer

THE ATTACHED PRINTOUT EXPLAINS DETECTED ERRORS.

PLEASE FORWARD THIS INFORMATION TO THE APPLICANT BY EITHER:

- 1) INCLUDING A COPY OF THIS PRINTOUT IN YOUR NEXT COMMUNICATION TO THE APPLICANT, WITH A NOTICE TO COMPLY or,
- 2) TELEPHONING APPLICANT AND FAXING A COPY OF THIS PRINTOUT, WITH A NOTICE TO COMPLY

FOR CRF SUBMISSION QUESTIONS, PLEASE CONTACT MARK SPENCER, 703-308-4212.

FOR SEQUENCE RULES INTERPRETATION, PLEASE CONTACT ROBERT WAX, 703-308-4216.

PATENTIN 2.1 e-mail help: [patin21help@uspto.gov](mailto:patin21help@uspto.gov) or phone 703-306-4119 (R. Wax)

PATENTIN 3.0 e-mail help: [patin30help@uspto.gov](mailto:patin30help@uspto.gov) or phone 703-306-4119 (R. Wax)

TO REDUCE ERRORED SEQUENCE LISTINGS, PLEASE USE THE CHECKER  
VERSION 3.0 PROGRAM, ACCESSIBLE THROUGH THE U.S. PATENT AND  
TRADEMARK OFFICE WEBSITE. SEE BELOW:

### Checker Version 3.0

The Checker Version 3.0 application is a state-of-the-art Windows based software program employing a logical and intuitive user-interface to check whether a sequence listing is in compliance with format and content rules. Checker Version 3.0 works for sequence listings generated for the original version of 37 CFR §§1.821 - 1.825 effective October 1, 1990 (old rules) and the revised version (new rules) effective July 1, 1998 as well as World Intellectual Property Organization (WIPO) Standard ST.25.

Checker Version 3.0 replaces the previous DOS-based version of Checker, and is Y2K-compliant. Checker allows public users to check sequence listings in Computer Readable form (CRF) before submitting them to the United States Patent and Trademark Office (USPTO). Use of Checker prior to filing the sequence listing is expected to result in fewer errored sequence listings, thus saving time and money.

Checker Version 3.0 can be down loaded from the USPTO website at the following address:

<http://www.uspto.gov/web/offices/pac/checker>

# Raw Sequence Listing Error Summary

## ERROR DETECTED SUGGESTED CORRECTION

SERIAL NUMBER:

09/284,697

ATTN: NEW RULES CASES: PLEASE DISREGARD ENGLISH "ALPHA" HEADERS, WHICH WERE INSERTED BY PTO SOFTWARE

- 1        Wrapped Nucleics      The number/text at the end of each line "wrapped" down to the next line.  
This may occur if your file was retrieved in a word processor after creating it.  
Please adjust your right margin to .3, as this will prevent "wrapping".
- 2        Wrapped Aminos      The amino acid-number/text at the end of each line "wrapped" down to the next line.  
This may occur if your file was retrieved in a word processor after creating it.  
Please adjust your right margin to .3, as this will prevent "wrapping".
- 3        Incorrect Line Length      The rules require that a line not exceed 72 characters in length. This includes spaces.
- 4        Misaligned Amino Acid      The numbering under each 5th amino acid is misaligned. This may be caused by the use of tabs  
Numbering      between the numbering. It is recommended to delete any tabs and use spacing between the numbers.
- 5   J   Non-ASCII      This file was not saved in ASCII (DOS) text, as required by the Sequence Rules.  
Please ensure your subsequent submission is saved in ASCII text so that it can be processed.
- 6        Variable Length      Sequence(s)        contain n's or Xaa's which represented more than one residue.  
As per the rules, each n or Xaa can only represent a single residue.  
Please present the maximum number of each residue having variable length and  
indicate in the (ix) feature section that some may be missing.
- 7        PatentIn ver. 2.0 "bug"      A "bug" in PatentIn version 2.0 has caused the <220>-<223> section to be missing from amino acid  
sequence(s)       . Normally, PatentIn would automatically generate this section from the  
previously coded nucleic acid sequence. Please manually copy the relevant <220>-<223> section  
to the subsequent amino acid sequence. This applies primarily to the mandatory <220>-<223>  
sections for Artificial or Unknown sequences.
- 8        Skipped Sequences      Sequence(s)        missing. If intentional, please use the following format for each skipped sequence:  
(OLD RULES)      (2) INFORMATION FOR SEQ ID NO:X:  
(i) SEQUENCE CHARACTERISTICS:(Do not insert any headings under "SEQUENCE CHARACTERISTICS")  
(xi) SEQUENCE DESCRIPTION:SEQ ID NO:X:  
This sequence is intentionally skipped  
  
Please also adjust the "(iii) NUMBER OF SEQUENCES:" response to include the skipped sequence(s).
- 9        Skipped Sequences      Sequence(s)        missing. If intentional, please use the following format for each skipped sequence.  
(NEW RULES)      <210> sequence id number  
                         <400> sequence id number  
                         000.
- 10        Use of n's or Xaa's      Use of n's and/or Xaa's have been detected in the Sequence Listing.  
(NEW RULES)      Use of <220> to <223> is MANDATORY if n's or Xaa's are present.  
In <220> to <223> section, please explain location of n or Xaa, and which residue n or Xaa represents.
- 11        Use of <213>Organism      Sequence(s)        are missing this mandatory field or its response.  
(NEW RULES)
- 12        Use of <220>Feature      Sequence(s)        are missing the <220>Feature and associated headings.  
(NEW RULES)      Use of <220> to <223> is MANDATORY if <213>ORGANISM is "Artificial" or "Unknown"  
                         Please explain source of genetic material in <220> to <223> section.  
                         (See "Federal Register," 6/01/98, Vol. 63, No. 104, pp. 29631-32) (Sec. 1.823 of new Rules)
- 13        PatentIn ver. 2.0 "bug"      Please do not use "Copy to Disk" function of PatentIn version 2.0. This causes a corrupted  
                         file, resulting in missing mandatory numeric identifiers and responses (as indicated on raw sequence listing).  
                         Instead, please use "File Manager" or any other means to copy file to floppy disk.

P:BU1

#10/12/19/20  
1638

**IMPORTANT:** see item 5 on  
Erra summary sheet  
Does Not Comply  
Corrected Diskette Needed

RAW SEQUENCE LISTING  
PATENT APPLICATION: US/09/284,697

DATE: 12/01/2000  
TIME: 08:23:23

Input Set : A:\ES.txt  
Output Set: N:\CRF3\12012000\I284697.raw

pp 1-5

OK

```
3 <110> APPLICANT: MERISTEM THERAPEUTICS S.A.
5 <120> TITLE OF INVENTION: Pancreatic Lipases and/or Recombinant Colipases and
6   Derived Polypeptides Produced by Plants, Methods for
7   Obtaining Them and Use thereof.
9 <130> FILE REFERENCE: 1074-1168PCF-US octobre 2000
11 <140> CURRENT APPLICATION NUMBER: US 09/284,697
12 <141> CURRENT FILING DATE: 1999-07-06
14 <150> PRIOR APPLICATION NUMBER: PCI/TR97/01862
15 <151> PRIOR FILING DATE: 1997-10-17
17 <160> NUMBER OF SEQ ID NOS: 16
19 <170> SOFTWARE: Patentin Ver. 2.1
21 <210> SEQ ID NO: 1
22 <211> LENGTH: 69
23 <212> TYPE: DNA
24 <213> ORGANISM: Artificial Sequence
26 <220> FEATURE:
27 <223> OTHER INFORMATION: Description of Artificial Sequence: Signal peptide
29 <220> FEATURE:
30 <221> NAME/KEY: misc_feature
31 <222> LOCATION: (1)..(69)
32 <223> OTHER INFORMATION: Signal peptide of Sporamine A of Sweet potato
34 <220> FEATURE:
35 <221> NAME/KEY: CDS
36 <222> LOCATION: (1)..(69)
38 <300> PUBLICATION INFORMATION:
39 <303> JOURNAL: J. Biol. Chem.
40 <304> VOLUME: 264
41 <306> PAGES: 20042-20048
42 <307> DATE: 1989
44 <400> SEQUENCE: 1
45 atg aaa gcc ttc aca ctc gct ctc ttc tta gct ctc tcc ctc tat ctc 48
46 Met Lys Ala Phe Thr Leu Ala Leu Phe Leu Ala Leu Ser Leu Tyr Leu
47 1 5 10 15
49 ctg ccc aat cca gcc cat tcc 69
50 Leu Pro Asn Pro Ala His Ser
51 20
54 <210> SEQ ID NO: 2
55 <211> LENGTH: 23
56 <212> TYPE: PRT
57 <213> ORGANISM: Artificial Sequence
W--> 58 <220> FEATURE: < insert this mandatory numeric identifier whenever
58 <223> OTHER INFORMATION: Description of Artificial Sequence: Signal peptide
60 <400> SEQUENCE: 2
61 Met Lys Ala Phe Thr Leu Ala Leu Phe Leu Ala Leu Ser Leu Tyr Leu
62 1 5 10 15
64 Leu Pro Asn Pro Ala His Ser
65 20
```

Sequence 1 is a nucleotide sequence.

This would be a sufficient  
Explanation for (213) Artificial  
Sequence.

give source of genetic material  
C2217, C2227, or  
C2237 is  
stout

see circled portion of item 12  
on Erra summary sheet

RAW SEQUENCE LISTING  
 PATENT APPLICATION: US/09/284,697  
 DATE: 12/01/2000  
 TIME: 08:23:23

Input Set : A:\ES.txt  
 Output Set: N:\CRF3\12012000\I284697.raw

69 <210> SEQ ID NO: 3  
 70 <211> LENGTH: 111  
 71 <212> TYPE: DNA  
 72 <213> ORGANISM: Artificial Sequence  
 74 <220> FEATURE:  
 75 <223> OTHER INFORMATION: Description of Artificial Sequence: Signal peptide  
 77 <220> FEATURE:  
 78 <221> NAME/KEY: sig\_peptide  
 79 <222> LOCATION: (1)..(111)  
 80 <223> OTHER INFORMATION: Signal peptide PPS of Sporamine A of Sweet potato.  
 82 <220> FEATURE:  
 83 <221> NAME/KEY: CDS  
 84 <222> LOCATION: (1)..(111)  
 86 <300> PUBLICATION INFORMATION:  
 87 <303> JOURNAL: J. Biol. Chem.  
 88 <304> VOLUME: 264  
 89 <306> PAGES: 20042-20048  
 90 <307> DATE: 1989

92 <400> SEQUENCE: 3  
 93 atg aaa gcc ttc aca ctc gct ctc tta gct ctt tcc ctc tat ctc 48  
 94 Met Lys Ala Phe Thr Leu Ala Leu Phe Leu Ala Leu Ser Leu Tyr Leu  
 95 1 5 10 15  
 97 ctg ccc aat cca gcc cat tcc agg ttc aat ccc atc cgc ctc ccc acc 96  
 98 Leu Pro Asn Pro Ala His Ser Arg Phe Asn Pro Ile Arg Leu Pro Thr  
 99 20 25 30  
 101 aca cac gaa ccc gcc 111  
 102 Thr His Glu Pro Ala  
 103 35

106 <210> SEQ ID NO: 4  
 107 <211> LENGTH: 37  
 108 <212> TYPE: PRI  
 109 <213> ORGANISM: Artificial Sequence  
 110 <220> FEATURE:  
 111 <223> OTHER INFORMATION: Description of Artificial Sequence: Signal peptide  
 112 <400> SEQUENCE: 4  
 113 Met Lys Ala Phe Thr Leu Ala Leu Phe Leu Ala Leu Ser Leu Tyr Leu  
 114 1 5 10 15  
 116 Leu Pro Asn Pro Ala His Ser Arg Phe Asn Pro Ile Arg Leu Pro Thr  
 117 20 25 30  
 119 Thr His Glu Pro Ala  
 120 35

124 <210> SEQ ID NO: 5  
 125 <211> LENGTH: 66  
 126 <212> TYPE: DNA  
 127 <213> ORGANISM: Artificial Sequence  
 129 <220> FEATURE:  
 130 <223> OTHER INFORMATION: Description of Artificial Sequence: Signal peptide  
 132 <220> FEATURE:  
 133 <221> NAME/KEY: sig\_peptide

*seq 3 is a nucleotide sequence*

*Signal peptide*

*Sweet*

*OK*

*insert*

*give source*

*Sequence 5 is a nucleotide sequence*



RAW SEQUENCE LISTING  
 PATENT APPLICATION: US/09/284,697

DATE: 12/01/2000  
 TIME: 09:23:23

Input Set : A:\ES.txt  
 Output Set: N:\CRF3\12012000\1284697.raw

134 <222> LOCATION: (1)..(66)  
 135 <223> OTHER INFORMATION: Signal peptide of rabbit gastric lipase  
 137 <220> FEATURE:  
 138 <221> NAME/KEY: CDS  
 139 <222> LOCATION: (1)..(66)  
 141 <400> SEQUENCE: 5  
 142 atg tgg qlg ctt ttc atg qtg qca gct tlg cta tct gca ctt gga act 48  
 143 Met Trp Val Leu Phe Met Val Ala Ala Leu Ser Ala Leu Gly Thr  
 144 1 5 10 15  
 146 aca cat ggt ctt ttt gga 66  
 147 Thr His Gly Leu Phe Gly  
 148 20

151 <210> SEQ ID NO: 6  
 152 <211> LENGTH: 22  
 153 <212> TYPE: PRI  
 154 <213> ORGANISM: Artificial Sequence

W--> 155 <220> FEATURE: *insert*

155 <223> OTHER INFORMATION: Description of Artificial Sequence Signal peptide  
 157 <400> SEQUENCE: 6  
 158 Met Trp Val Leu Phe Met Val Ala Ala Leu Ser Ala Leu Gly Thr  
 159 1 5 10 15  
 161 Thr His Gly Leu Phe Gly  
 162 20

166 <210> SEQ ID NO: 7  
 167 <211> LENGTH: 48  
 168 <212> TYPE: DNA  
 169 <213> ORGANISM: Artificial Sequence

171 <220> FEATURE:  
 172 <223> OTHER INFORMATION: Description of Artificial Sequence Signal peptide  
 174 <220> FEATURE:  
 175 <221> NAME/KEY: sig\_peptide

176 <222> LOCATION: (1)..(48)  
 177 <223> OTHER INFORMATION: Signal peptide of PSHPL  
 179 <220> FEATURE:

180 <221> NAME/KEY: CDS  
 181 <222> LOCATION: (1)..(48)  
 182 <223> OTHER INFORMATION: The cleavage sequence between the two sequences  
 183 coding for PSHPL and NPL is Gly-Lys.

185 <400> SEQUENCE: 7  
 186 atg ctg cca ctt tgg act ctt tca ctg ctg ctg gga gca gla gca gga 48  
 187 Met Leu Pro Leu Trp Thr Leu Ser Leu Leu Gly Ala Val Ala Gly  
 188 1 5 10 15

191 <210> SEQ ID NO: 8  
 192 <211> LENGTH: 16  
 193 <212> TYPE: PRI  
 194 <213> ORGANISM: Artificial Sequence

W--> 195 <220> FEATURE: *insert*

195 <223> OTHER INFORMATION: Description of Artificial Sequence Signal peptide  
 197 <400> SEQUENCE: 8

*do you mean rabbit?*

*give source*

*nucleotide sequence*

*give source*

RAW SEQUENCE LISTING  
PATENT APPLICATION: US/09/284,697

DATE: 12/01/2000  
TIME: 08:23:23

Input Set : A:\ES.txt  
Output Set: N:\CRF3\12012000\I284697.raw

198 Met Leu Pro Leu Trp Thr Leu Ser Leu Leu Leu Gly Ala Val Ala Gly  
199 1 5 10 15  
203 <210> SEQ ID NO: 9  
204 <211> LENGTH: 66  
205 <212> TYPE: DNA  
206 <213> ORGANISM: Artificial Sequence  
208 <220> FEATURE:  
209 <223> OTHER INFORMATION: Description of Artificial Sequence: Signal peptide  
211 <220> FEATURE:  
212 <221> NAME/KEY: sig\_peptide  
213 <222> LOCATION: (1)..(66)  
214 <223> OTHER INFORMATION: Signal peptide of HPCOL  
216 <220> FEATURE:  
217 <221> NAME/KEY: CDS  
218 <222> LOCATION: (1)..(66)  
220 <400> SEQUENCE: 9  
221 atg tgg atg ctt ttc atg atg gca gct ttg cta tct gca ctt gga act 48  
222 Met Trp Val Leu Phe Met Val Ala Ala Leu Leu Ser Ala Leu Gly Thr  
223 1 5 10 15  
225 aca cat ggt ctt ttt gga 66  
226 Thr His Gly Leu Phe Gly  
227 20  
230 <210> SEQ ID NO: 10  
231 <211> LENGTH: 22  
232 <212> TYPE: DRT  
233 <213> ORGANISM: Artificial Sequence  
W--> 234 <220> FEATURE: insert  
234 <223> OTHER INFORMATION: Description of Artificial Sequence: Signal peptide  
236 <400> SEQUENCE: 10  
237 Met Trp Val Leu Phe Met Val Ala Ala Leu Leu Ser Ala Leu Gly Thr  
238 1 5 10 15  
240 Thr His Gly Leu Phe Gly  
241 20  
245 <210> SEQ ID NO: 11  
246 <211> LENGTH: 51  
247 <212> TYPE: DNA  
248 <213> ORGANISM: Artificial Sequence  
250 <220> FEATURE:  
251 <223> OTHER INFORMATION: Description of Artificial Sequence: Signal peptide  
253 <220> FEATURE:  
254 <221> NAME/KEY: sig\_peptide  
255 <222> LOCATION: (1)..(51)  
256 <223> OTHER INFORMATION: Signal peptide of PSHPOL  
258 <220> FEATURE:  
259 <221> NAME/KEY: CDS  
260 <222> LOCATION: (1)..(51)  
261 <223> OTHER INFORMATION: The cleavage sequence between the two sequences  
262 coding for PSHPOL and HPCOL is Ala-Lys.  
264 <400> SEQUENCE: 11

*nucleotide sequence*

*gene source*

*nucleotide sequence*

VERIFICATION SUMMARY  
PATENT APPLICATION: US/09/284,697  
DATE: 12/01/2000  
TIME: 08:23:24

Input Set : A:\ES.txt  
Output Set: N:\CRF3\12012000\I284697.raw

L:12 M:271 C: Current Filing Date differs. Replaced Current Filing Date  
L:42 M:285 W: Invalid Journal Date: Wrong YYYY-MM-DD,MMM-YYYY or SEASON-YYYY. Wrong Journal Date:YYYY-MM-DD,MMM-YYYY or Season-YYYY  
L:58 M:258 W: Mandatory Feature missing, <220> FEATURE:  
L:90 M:285 W: Invalid Journal Date: Wrong YYYY-MM-DD,MMM-YYYY or SEASON-YYYY. Wrong Journal Date:YYYY-MM-DD,MMM-YYYY or Season-YYYY  
L:110 M:258 W: Mandatory Feature missing, <220> FEATURE:  
L:155 M:258 W: Mandatory Feature missing, <220> FEATURE:  
L:195 M:258 W: Mandatory Feature missing, <220> FEATURE:  
L:234 M:258 W: Mandatory Feature missing, <220> FEATURE:  
L:277 M:258 W: Mandatory Feature missing, <220> FEATURE: